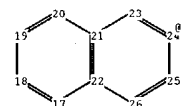
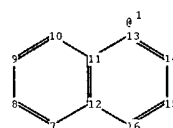
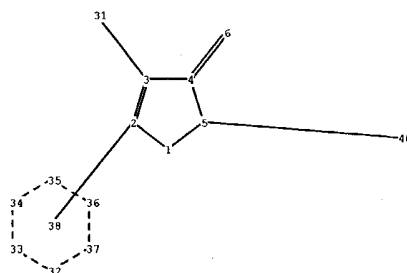
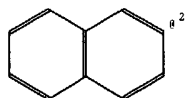
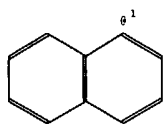
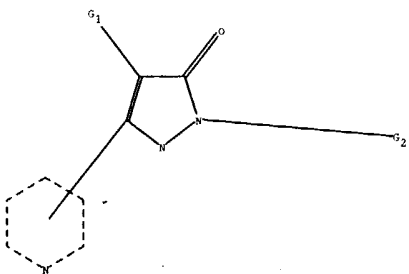


C:\stnweb\Queries\7.str



chain nodes :

6 31 40

ring nodes :

1 2 3 4 5 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25
26 32 33 34 35 36 37

chain bonds :

3-31 4-6 5-40

ring bonds :

1-2 1-5 2-3 3-4 4-5 7-8 7-12 8-9 9-10 10-11 11-12 11-13 12-16 13-14 14-15
15-16 17-18 17-22 18-19 19-20 20-21 21-22 21-23 22-26 23-24 24-25 25-26 32-33
32-37 33-34 34-35 35-36 36-37

exact/norm bonds :

1-2 1-5 3-31 4-5 4-6 5-40 32-33 32-37 33-34 34-35 35-36 36-37

exact bonds :

2-3 3-4

normalized bonds :

7-8 7-12 8-9 9-10 10-11 11-12 11-13 12-16 13-14 14-15 15-16 17-18 17-22
18-19 19-20 20-21 21-22 21-23 22-26 23-24 24-25 25-26

isolated ring systems :

containing 1 : 7 : 17 :

G1:Ph,[*1],[*2]

G2:H,Ak

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:CLASS 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom
12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom
22:Atom 23:Atom 24:Atom 25:Atom 26:Atom 31:CLASS 32:Atom 33:Atom 34:Atom 35:Atom
36:Atom 37:Atom 38:CLASS 40:CLASS

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 NEWS 2 "Ask CAS" for self-help around the clock
 NEWS 3 Jul 12 BEILSTEIN enhanced with new display and select options,
 resulting in a closer connection to BABS
 NEWS 4 Jul 30 BEILSTEIN on STN workshop to be held August 24 in conjunction
 with the 228th ACS National Meeting
 NEWS 5 AUG 02 IFIPAT/IFIUDB/IFICDB reloaded with new search and display
 fields
 NEWS 6 AUG 02 CAPLUS and CA patent records enhanced with European and Japan
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 NEWS 7 AUG 02 The Analysis Edition of STN Express with Discover!
 (Version 7.01 for Windows) now available
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 NEWS 9 AUG 27 BIOCOMMERCE: Changes and enhancements to content coverage
 NEWS 10 AUG 27 BIOTECHABS/BIOTECHDS: Two new display fields added for legal
 status data from INPADOC
 NEWS 11 SEP 01 INPADOC: New family current-awareness alert (SDI) available
 NEWS 12 SEP 01 New pricing for the Save Answers for SciFinder Wizard within
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 NEWS 13 SEP 01 New display format, HITSTR, available in WPIDS/WPINDEX/WPIX
 NEWS 14 SEP 14 STN Patent Forum to be held October 13, 2004, in Iselin, NJ
 NEWS 15 SEP 27 STANDARDS will no longer be available on STN
 NEWS 16 SEP 27 SWETSCAN will no longer be available on STN

NEWS EXPRESS JULY 30 CURRENT WINDOWS VERSION IS V7.01, CURRENT
 MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
 AND CURRENT DISCOVER FILE IS DATED 11 AUGUST 2004

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=> file reg

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FULL ESTIMATED COST	0.21	0.21

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STRUCTURE FILE UPDATES: 26 SEP 2004 HIGHEST RN 752189-88-1

DICTIONARY FILE UPDATES: 26 SEP 2004 HIGHEST RN 752189-88-1

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

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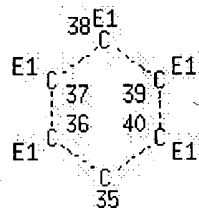
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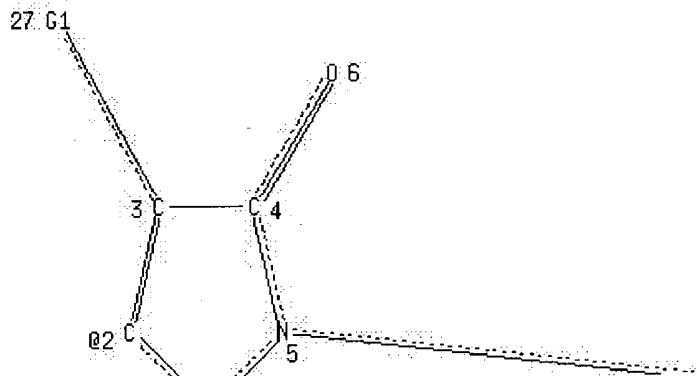
L1 HAS NO ANSWERS

L1 STR

H 41 Ak 42



Page 1-A



Page 1-B

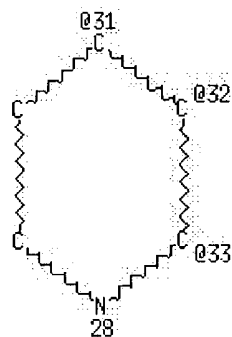
030

029

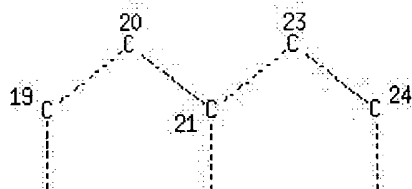
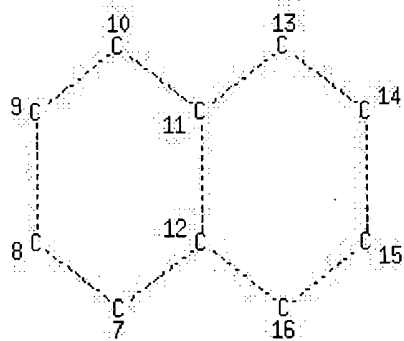
Page 2-A



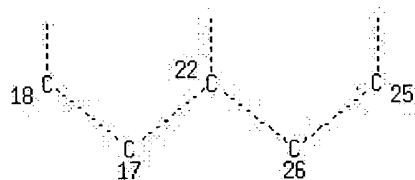
G2 34



Page 2-B



Page 3-B



Page 4-B

VAR G1=35/13/24

VAR G2=41/42

VPA 2-29/30/31/32/33 S

NODE ATTRIBUTES:

HCOUNT	IS	E1	AT	36
HCOUNT	IS	E1	AT	37
HCOUNT	IS	E1	AT	38
HCOUNT	IS	E1	AT	39
HCOUNT	IS	E1	AT	40
NSPEC	IS	R	AT	1
NSPEC	IS	R	AT	2
NSPEC	IS	R	AT	3
NSPEC	IS	R	AT	4
NSPEC	IS	R	AT	5

NSPEC IS C AT 6
 NSPEC IS R AT 7
 NSPEC IS R AT 8
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 NSPEC IS C AT 27
 NSPEC IS R AT 28
 NSPEC IS R AT 29
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 NSPEC IS R AT 32
 NSPEC IS R AT 33
 NSPEC IS C AT 34
 DEFAULT MLEVEL IS ATOM
 MLEVEL IS CLASS AT 6 35 36 37 38 39 40 41 42
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RSPEC 1 7 17
 NUMBER OF NODES IS 42

STEREO ATTRIBUTES: NONE

=> s 11

SAMPLE SEARCH INITIATED 14:31:29 FILE 'REGISTRY'
 SAMPLE SCREEN SEARCH COMPLETED - 434 TO ITERATE

100.0% PROCESSED 434 ITERATIONS 0 ANSWERS
 SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
 BATCH **COMPLETE**
 PROJECTED ITERATIONS: 7431 TO 9929
 PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s 11 full

THE ESTIMATED SEARCH COST FOR FILE 'REGISTRY' IS 155.00 U.S. DOLLARS
 DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N or END:y
 FULL SEARCH INITIATED 14:31:34 FILE 'REGISTRY'
 FULL SCREEN SEARCH COMPLETED - 9246 TO ITERATE

100.0% PROCESSED 9246 ITERATIONS
SEARCH TIME: 00.00.01

3 ANSWERS

L3 3 SEA SSS FUL L1

=> file hcaplus

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

162.98

163.19

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FILE COVERS 1907 - 27 Sep 2004 VOL 141 ISS 14
FILE LAST UPDATED: 26 Sep 2004 (20040926/ED)

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=> s 13

L4 1 L3

=> d 14, ibib abs fhitstr, 1

L4 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN

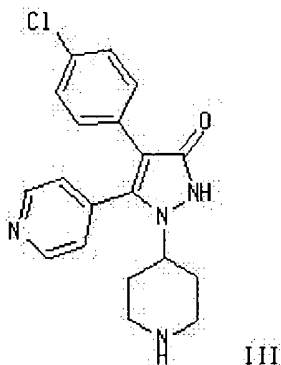
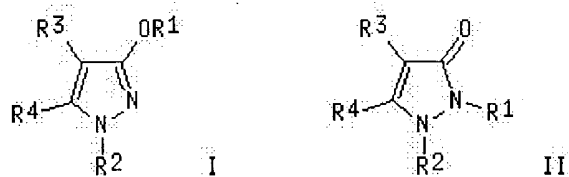
Full Text References

ACCESSION NUMBER: 2004:220202 HCAPLUS
DOCUMENT NUMBER: 140:253561
TITLE: Preparation of 1,2-dihydropyrazol-3-ones and 3-alkoxy-1H-pyrazoles as TNF- α and interleukin lowering agents for the treatment of inflammation
INVENTOR(S): Dominguez, Celia; Zhang, Dawei; Sham, Kelvin K. C.; Cao, Guo-qiang
PATENT ASSIGNEE(S): Amgen Inc., USA
SOURCE: PCT Int. Appl., 63 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004022055	A1	20040318	WO 2003-US28067	20030908
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,				

CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
 GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
 LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
 PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA,
 UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
 CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC,
 NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ,
 GW, ML, MR, NE, SN, TD, TG

US 2004058918 A1 20040325 US 2003-658298 20030908
 PRIORITY APPLN. INFO.: US 2002-409176P P 20020909
 OTHER SOURCE(S): MARPAT 140:253561
 GI



AB The invention discloses the prepn. of title compds. I and II [wherein R1 = H or alkyl; R2 = alkyl, Ph, PhCH₂, (alkyl)R_c, (alkyl)R_f, or R_g; R3 and R4 = independently (un)substituted Ph, naphthyl, or heterocyclyl; R_c = independently (un)substituted heterocyclyl; R_f = substituted R_c; R_g = substituted alkyl, Ph, or PhCH₂; and pharmaceutically acceptable salts thereof] as tumor necrosis factor α (TNF α) and interleukin 1, 6, and 8 (IL-1, IL-6, and IL-8) inhibitors. For example, (4-chlorophenyl)acetic acid was condensed with pyridine-4-carbaldehyde in acetic anhydride and TEA to give 2-(4-chlorophenyl)-3-(pyridin-4-yl)acrylic acid, which was esterified with MeOH in thionyl chloride. Cyclization of the acrylate with hydrazine in EtOH, followed by Pd/C catalyzed redn., afforded 4-(4-chlorophenyl)-5-(pyridin-4-yl)-1,2-dihydropyrazol-3-one. Addn. of 4-oxopiperidine-1-carboxylic acid tert-Bu ester in chloroform using sodium triacetoxy boron hydride, hydrogenation using Pd/C in EtOH, and deprotection with HCl in ether and dioxane gave III. Selected compds. of the invention inhibited lipopolysaccharide-activated TNF prodn. in THP1 cells with IC₅₀ values of <20 μ M. Also disclosed is a method of prophylaxis or treatment of inflammation, rheumatoid arthritis, Paget's disease, osteoporosis, multiple myeloma, uveitis, acute or chronic myelogenous leukemia, pancreatic β cell destruction, osteoarthritis, rheumatoid spondylitis, gouty arthritis,

inflammatory bowel disease, adult respiratory distress syndrome (ARDS), psoriasis, Crohn's disease, allergic rhinitis, ulcerative colitis, anaphylaxis, contact dermatitis, asthma, muscle degeneration, cachexia, Reiter's syndrome, type I diabetes, type II diabetes, bone resorption diseases, graft vs. host reaction, Alzheimer's disease, stroke, myocardial infarction, ischemia reperfusion injury, atherosclerosis, brain trauma, multiple sclerosis, cerebral malaria, sepsis, septic shock, toxic shock syndrome, fever, myalgias due to HIV-1, HIV-2, HIV-3, cytomegalovirus (CMV), influenza, adenovirus, the herpes viruses, or herpes zoster infection in a mammal comprising administering an effective amt. of I or II or their pharmaceutical compns. (no data).

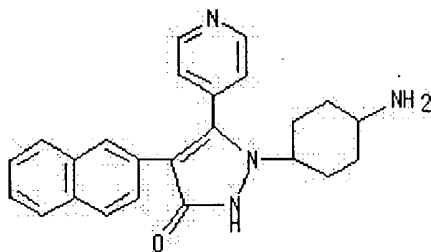
IT 671781-23-0P, 1-(4-Aminocyclohexyl)-4-(naphthalen-2-yl)-5-(pyridin-4-yl)-1,2-dihydropyrazol-3-one

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(TNF α and/or IL inhibitor; prepn. of dihydropyrazolones and alkoxy pyrazoles as TNF- α and interleukin lowering agents for treatment of inflammation and related conditions)

RN 671781-23-0 HCAPLUS

CN 3H-Pyrazol-3-one, 1-(4-aminocyclohexyl)-1,2-dihydro-4-(2-naphthalenyl)-5-(4-pyridinyl)- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> file caold

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
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FILE COVERS 1907-1966

FILE LAST UPDATED: 01 May 1997 (19970501/UP)

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L1 STRUCTURE UPLOADED

L2 0 S L1

L3 3 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 14:31:37 ON 27 SEP 2004

L4 1 S L3

FILE 'CAOLD' ENTERED AT 14:31:47 ON 27 SEP 2004

=> s l3

L5 0 L3

=>